Co

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

in the application of: Blake Pepinsky et al.

Docket No.: 14937.0059

Filed: March 16, 2004

DEC 1 7 2008

Issued: November 4, 2008

Serial No.: 10/802,540

Patent No.: 7,446,173 B2

For: POLYMER CONJUGATES OF INTERFERON BETA-1A AND USES

ATTN: Certificate of Correction Branch United States Patent and Trademark Office Customer Service Window Randolph Building 401 Dulany Street Alexandria, VA 22314

Certificate

DEC 1 9 2008

of Correction

REQUEST FOR EXPEDITED ISSUANCE OF CERTIFICATE OF CORRECTION PURSUANT TO 37 C.F.R. 1.322

Applicants respectfully request that a Certificate of Correction be issued to correct the omission of SEQ ID NOs: 41-56 of the above-mentioned patent. The omission of SEQ ID NOs: 41-56 was incurred by the U.S. Patent and Trademark Office. A Supplemental Preliminary Amendment filed on August 18, 2006 provided a substitute sequence listing which included SEQ ID NOs: 41-56. The substitute sequence listing was entered by the USPTO on August 22, 2006. Applicants herein submit a copy of the filed Supplemental Preliminary Amendment as Exhibit A. A copy of the Entered Raw Sequence Listing is provided as Exhibit B. A Certificate of Correction form, PTO/SB/44 is also submitted herewith.

Applicants do not believe that any fees are due with the filing as the error in the claims was incurred by the USPTO. However, should any fees be required by this request, the Commissioner is hereby authorized to charge Deposit Account 19-4293.

Respectfully submitted,

Date: 12-17-08

Harold H. Fox

Reg. No. 41,498

Steptoe & Johnson LLP 1330 Connecticut Avenue, NW Washington, DC 20036-1795

Phone: 202-429-3000 Fax: 202-429-3902

DEC 1 9 2008

PATENT NO.

: 7,446,173 B2

APPLICATION NO.: 10/802,540

ISSUE DATE

: November 4, 2008

INVENTOR(S)

: PEPINSKY ET AL.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

Column 63, line 35, insert the following SEQ ID NOs: 41-56:

<210> 41

<211> 166

<212> PRT

<213> Homo sapiens

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 135

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu 150

Thr Gly Tyr Leu Arg Asn

165

MAILING ADDRESS OF SENDER:

PATENT No. 7,446,173 B2

PATENT No. : 7,446,173 B2

APPLICATION NO. : 10/802,540

ISSUE DATE: NOVEMBER 4, 2008

Inventor(S) : PEPINSKY ET AL.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 42

<211> 166

<212> PRT <213> Homo sapiens

<400> 42

Met Ala Tyr Ala Ala Leu Gly Ala Leu Gln Ala Ser Ser Asn Phe Gln 5 10 . 15

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 . 110

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 120

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 135

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu

Thr Gly Tyr Leu Arg Asn

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PATENT No. 7,446,173 B2

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APPLICATION NO.: 10/802,540

ISSUE DATE : NOVEMBER 4, 2008

INVENTOR(S) : PEPINSKY ET AL.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 43

<211> 166

<212> PRT

<213> Homo sapiens

<400> 43

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Ala Ala 1 5 10 15

Cys Ala Ala Leu Leu Ala Ala Leu Asn Gly Arg Leu Glu Tyr Cys Leu 20 25 30

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu 145 150 155 160

Thr Gly Tyr Leu Arg Asn 165

MAILING ADDRESS OF SENDER:

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ISSUE DATE: NOVEMBER 4, 2008

INVENTOR(S) : PEPINSKY ET AL.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 44

<211> 166 <212> PRT

<213> Homo sapiens

<400> 44

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Ala Ala Ala Cys Ala 20 25 30

Ala Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu 145 150 155 160

Thr Gly Tyr Leu Arg Asn 165

MAILING ADDRESS OF SENDER:

PATENT No. 7,446,173 B2

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APPLICATION NO. : 10/802,540

ISSUE DATE: NOVEMBER 4, 2008

INVENTOR(S) : PEPINSKY ET AL.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 45

<211> 166

<212> PRT

<213> Homo sapiens

<400> 45

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln
1 5 10 15

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu 20 25 30

Lys Asp Arg Ala Ala Phe Ala Ile Pro Ala Giu Ile Lys Gin Leu Gin 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 $$ 55 $$ 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu 145 150 155 160

Thr Gly Tyr Leu Arg Asn 165

MAILING ADDRESS OF SENDER:

PATENT No. 7,446,173 B2

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APPLICATION NO.: 10/802,540

: November 4, 2008 ISSUE DATE

Inventor(S) : PEPINSKY ET AL.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 46

<211> 166

<212> PRT

<213> Homo sapiens

<400> 46

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Ala Ala Ala Ala

Ala Phe Ala Ala Ala Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 55

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 135

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu

Thr Gly Tyr Leu Arg Asn 165

MAILING ADDRESS OF SENDER:

PATENT No. 7,446,173 B2

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APPLICATION NO. : 10/802,540

ISSUE DATE : November 4, 2008

INVENTOR(S) : PEPINSKY ET AL.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 47

<211> 166

<212> PRT <213> Homo sapiens

<400> 47

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Ala

Asn Ile Ala Ser Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 135

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu

Thr Gly Tyr Leu Arg Asn

MAILING ADDRESS OF SENDER:

PATENT No. _____7,446,173 B2

: 7,446,173 B2 PATENT NO.

APPLICATION NO. : 10/802,540

: November 4, 2008 ISSUE DATE

: PEPINSKY ET AL. Inventor(S)

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 48

<211> 166

<212> PRT

<213> Homo sapiens

<400> 48

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu 20 30 .

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln

Asn Ile Phe Ala Ile Phe Ala Ala Ala Ser Ser Ser Thr Gly Trp Asn

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 135

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu 145 150 155

Thr Gly Tyr Leu Arg Asn

MAILING ADDRESS OF SENDER:

PATENT No. 7,446,173 B2

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APPLICATION NO.: 10/802,540

ISSUE DATE : NOVEMBER 4, 2008

INVENTOR(S) : PEPINSKY ET AL.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 49

<211> 166
<212> PRT

<213> Homo sapiens

<400> 49

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln
1 5 10 15

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu 20 25 30

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Ala Ser Ile Val Ala Ala Leu Leu Ser Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu

Thr Gly Tyr Leu Arg Asn 165

MAILING ADDRESS OF SENDER:

PATENT No. 7,446,173 B2

PATENT NO. : 7,446,173 B2

APPLICATION NO. : 10/802,540

: November 4, 2008 ISSUE DATE

Inventor(S) : Pepinsky et al.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 50

<211> 166

<212> PRT

<213> Homo sapiens

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu 20 25 30

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 40

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Ala His Gln Ile Ala

His Leu Ala Ala Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 120

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu

Thr Gly Tyr Leu Arg Asn

MAILING ADDRESS OF SENDER:

PATENT No. 7,446,173 B2

United States Patent and Trademark Office **CERTIFICATE OF CORRECTION**

PATENT NO.

: 7,446,173 B2

APPLICATION NO. : 10/802,540

ISSUE DATE: NOVEMBER 4, 2008

Inventor(S)

: PEPINSKY ET AL.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 51

<211> 166

<212> PRT

<213> Homo sapiens

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu 20 25 30

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45 35 40

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn

His Leu Lys Thr Val Leu Ala Ala Lys Leu Ala Ala Ala Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 135

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu 150

Thr Gly Tyr Leu Arg Asn

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PATENT No. 7,446,173 B2

PATENT NO. : 7,446,173 B2

APPLICATION NO.: 10/802,540

ISSUE DATE : November 4, 2008

INVENTOR(S) : PEPINSKY ET AL.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 52

<211> 166 <212> PRT

<213> Homo sapiens

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu 20 25 30

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Ala Ala Thr 100 105

Ala Gly Lys Ala Met Ser Ala Leu His Leu Lys Arg Tyr Tyr Gly Arg

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu

Thr Gly Tyr Leu Arg Asn

165

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ISSUE DATE: NOVEMBER 4, 2008

INVENTOR(S) : PEPINSKY ET AL.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 53 <211> 166

<212> PRT

<213> Homo sapiens

<400> 53

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln
1 5 10 15

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu 20 25 30

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80.

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Ala 115 120 125

Ile Ala Ala Tyr Leu Ala Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu 145 150 155 160

Thr Gly Tyr Leu Arg Asn

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ISSUE DATE

: November 4, 2008

INVENTOR(S) : PEPINSKY ET AL.

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

<210> 54

<211> 166

<212> PRT

<213> Homo sapiens

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 40

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

Ile Leu His Tyr Leu Lys Ala Ala Ala Tyr Ser His Cys Ala Trp Thr 135

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu 150

Thr Gly Tyr Leu Arg Asn

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<210> 55

<211> 166

<212> PRT

<213> Homo sapiens

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 120

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ala Ala Cys Ala Trp Thr

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu 150

Thr Gly Tyr Leu Arg Asn

MAILING ADDRESS OF SENDER:

PATENT No. 7,446,173 B2

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APPLICATION NO.: 10/802,540

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<210> 56

<211> 166

<212> PRT

<213> Homo sapiens

<400> 56

Met Ser Tyr Asn Leu Leu Gly Phe Leu Gln Arg Ser Ser Asn Phe Gln
1 5 10 15

Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu 20 25 30

Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140

Ile Val Arg Ala Glu Ile Leu Ala Asn Phe Ala Phe Ile Ala Arg Leu 145 150 155 160

Thr Gly Tyr Leu Arg Asn 165

MAILING ADDRESS OF SENDER:

PATENT No. 7,446,173 B2





IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of: Pepinsky et al.

Serial No:

10/802,540

Filed:

March 16, 2004

For:

Polymer Conjugates of Interferon Beta-la

and Uses

Examiner:

Not yet known

Art Unit:

1646

Confirmation No.: 4023

Atty Docket No.: BII-008.02

CERTIFICATE OF FIRST CLASS MAILING

I hereby certify that this correspondence is being deposited with the United States Postal Service as First Class Mail, in an envelope addressed to Mail Stop Amendment, Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450, on August 16, 2006.

Merlin Aubourg

Mail Stop Amendment Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

SUPPLEMENTAL PRELIMINARY AMENDMENT

Dear Sir:

Prior to substantive examination of the above-referenced patent application, please amend the application as follows:

In the Claims:

1-40. (canceled)

- 41. (currently amended) A composition comprising a glycosylated interferon-beta-la comprising the amino acid sequence set forth in any one of SEQ ID NOs: 27-[[40]]56 coupled to a non-naturally-occurring polymer at an N-terminal end of said glycosylated interferon-beta-la, said polymer comprising a polyalkylene glycol moiety.
- 42. (previously presented) The composition of claim 41, wherein the polyalkylene moiety is coupled to the interferon-beta by way of a group selected from an aldehyde group, a maleimide group, a vinylsulfone group, a haloacetate group, plurality of histidine residues, a hydrazine group and an aminothiol group.
- 43. (currently amended) The composition of claim 41, wherein the interferon-beta-1a of any one of SEQ ID NOs: 27-[[40]]56 is an interferon-beta-1a fusion protein.
- 44. (previously presented) The composition of claim 43, wherein the interferon-beta-la fusion protein comprises a portion of an immunoglobulin molecule.
- 45. (currently amended) A physiologically active interferon-beta composition comprising a physiologically active interferon-beta-1a comprising an amino acid sequence selected from the group consisting of SEQ ID NOs: 27-[[40]]56, coupled to a polymer comprising a polyalkylene glycol moiety, wherein the interferon -beta-1a is coupled to the polymer at a site on the interferon-beta-1a that is an N- terminal end, wherein the physiologically active interferon -beta 1a and the polyalkylene glycol moiety are arranged such that the physiologically active interferon-beta-1a in the physiologically active interferon -beta composition has an activity at least 2-fold greater relative to physiologically active interferon-beta-1b, when measured by an antiviral assay.
- 46. (previously presented) The composition of claim 45, wherein the interferon-beta-1a is coupled to the polymer at a site by way of a glycan moiety of the interferon-beta-1a.
- 47. (previously presented) The composition of claim 45, wherein the interferon-beta-la is an interferon-beta-la fusion protein.
- 48. (previously presented) The composition of claim 47, wherein the interferon-beta-1a fusion protein comprises a portion of an immunoglobulin molecule.
- 49. (currently amended) A physiologically active interferon-beta composition comprising a physiologically active glycosylated interferon-beta-la comprising an amino acid sequence

selected from the group consisting of SEQ ID NO: 27-[[40]]56, N-terminally coupled to a polymer comprising a polyalkylene glycol moiety, wherein the physiologically active interferon-beta-1a and the polyalkylene glycol moiety are arranged such that the physiologically active interferon-beta 1a in the physiologically active interferon-beta composition has equal activity relative to physiologically active interferon-beta lacking said moiety, when measured by an antiviral assay.

. . . M

- 50. (previously presented) The composition of claim 49, wherein the interferon-beta is coupled to the polymer at a site by way of a glycan moiety on the interferon-beta.
- 51. (previously presented) The composition of claim 49, wherein the interferon-beta-la is an interferon beta fusion protein.
- 52. (previously presented) The composition of claim 51, wherein the interferon beta fusion protein comprises a portion of an immunoglobulin molecule.
- comprising a interferon-beta-la comprising an amino acid sequence selected from the group consisting of SEQ ID NOs: 27-[[40]]56, N-terminally coupled to a polyethylene glycol moiety, wherein the interferon-beta-la is coupled to the polyethylene glycol moiety by a labile bond, wherein the labile bond is cleavable by biochemical hydrolysis and/or protoclysis.
- 54. (previously presented) An interferon-beta composition according to claims 41, wherein the polymer has a molecular weight of from about 5 to 40 kilodaltons.
- 55. (previously presented) An interferon-beta composition according to claims 49, wherein the polymer has a molecular weight of from about 5 to 40 kilodaltons.
- 56. (previously presented) A interferon-beta composition according to claims 53, wherein the polymer has a molecular weight of from about 5 to 40 kilodaltons.
- 57. (previously presented) A pharmaceutical composition comprising the interferon-beta composition of claim 54.
- 58. (currently amended) A protein comprising the amino acid sequence set forth in any one of SEQ ID NOs: 25-[[40]]56 coupled to a non-naturally-occurring polymer at the C-terminal end of said protein, said polymer comprising a polyalkylene glycol moiety.
- 59. (currently amended) A protein comprising the amino acid sequence set forth in any one of SEQ ID NOs: 25-[[40]]56 coupled to a non-naturally-occurring polymer, said polymer comprising a

B3224018.1 - 3 -

- polyalkylene glycol moiety, and said polymer is attached to an amino, carboxylic, hydroxyl, guanidyl, or glycan moiety of said protein.
- 60. (currently amended) A protein comprising the amino acid sequence set forth in any one of SEQ ID NOs: 25-[[40]]56 coupled to a non-naturally-occurring polymer at the N-terminal end of said protein, said polymer comprising a polyalkylene glycol moiety.
- 61. (currently amended) A method of treating multiple sclerosis in a subject comprising administering to a subject in need thereof a therapeutically effect amount of a protein comprising the amino acid sequence set forth in any one of SEQ ID NOs: 25-[[40]]56 coupled to a non-naturally-occurring polymer, said polymer comprising a polyalkylene glycol moiety.
- 62. (currently amended) A method of preparing the protein of claim 60, comprising reacting a protein with a non-naturally-occurring polymer under reductive alkylation conditions, said protein comprising the amino acid sequence set forth in any one of SEQ ID NOs: 25-[[40]]56, and said polymer comprising a polyalkylene glycol moiety and a terminal aldehyde moiety.

Remarks

Claims 41-62 are pending. Claims 41, 43, 45, 49, and 53 were amended to refer to SEQ ID NOs: 27-56. Claims 58-62 were amended to refer to SEQ ID NOs: 25-56. A substitute Sequence Listing is submitted concurrently herewith. SEQ ID NOs: 1-40 of the Sequence Listing submitted herewith are the same as those in the paper copy of the sequence listing filed on July 19, 2004. SEQ ID NO: 41 is the amino acid sequence of wild-type IFN-beta-1a, which sequence is provided in Figure 10 of the application as the sequence spanning positions Meth18-Asn183. As stated on page 9 in the brief description for Figure 10, the amino acid sequence of wild-type IFN-beta-1a corresponds to the amino acid sequence spanning positions Meth18-Asn183 of the amino acid sequence displayed in Figure 10. SEQ ID NOs: 42-56 are the amino acid sequences of interferon-beta-1a mutants A1, A2, AB1, AB2, AB3, B1, B2, C1, C2, CD1, CD2, D, DE1, DE2, and E. As described on page 16 in lines 19-29 and further on pages 28-30 of the application, the amino acid sequences of the aforementioned interferon-beta-1a mutants are the same as that of wild-type IFN-beta-la except for certain alanine and/or serine mutations of the wild-type IFN-beta-1a amino acid sequence. The location of the alanine and/or serine mutations in the amino acid sequence for each interferon-beta-1a mutant is shown in Table 1 on page 32 of the application. Importantly, support for the claim amendments and the Sequence Listing submitted herewith can be found in the application. Therefore, no new matter has been added.

<u>Fees</u>

Applicants hereby authorize the Director to charge any required fee to our Deposit Account, No. 06-1448.

CONCLUSION

In view of the foregoing remarks, early and favorable consideration is respectfully solicited. The Examiner may address any questions raised by this submission to the undersigned at 617-832-1000.

155 Seaport Boulevard Boston, MA 02210

Telephone: (617) 832-1000 Telecopier: (617) 832-7000

Date: 8/16/06

Respectfully submitted,

Foley Hoag LLP

By:

Isabelle M. Clauss, Ph.D.

Reg. No. 47,326

Attorney for Applicants

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

भ्री re Application of: Pepinsky, et al.

Serial No:

10/802,540

Filed:

March 16, 2004

For:

Polymer Conjugates of Interferon Beta-

1A and Uses

Examiner:

Not Yet Assigned

Art Unit:

1646

Confirmation No.: 4023

Atty Docket No.: BII-008.02

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Merlin Aubourg

Mail Stop Amendment Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

STATEMENT UNDER 37 C.F.R. § 1.821(e), (f), and (g)

Sir:

In connection with a Sequence Listing submitted concurrently herewith, the undersigned hereby states that:

- 1. the submission, filed herewith in accordance with 37 C.F.R. § 1.821(g), does not include new matter;
- 2. the content of the attached paper copy and the attached computer readable copy of the Sequence Listing, submitted in accordance with 37 C.F.R. § 1.821(c) and (e), respectively, are the same.

Respectfully submitted, Foley Hoag LLP

By:

Chad E. Davis, Ph.D. Reg. No. 56,179

Agent for Applicants

155 Seaport Boulevard Boston, MA 02210

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Date:

8/16/06



SEQUENCE LISTING

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Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95 His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Ala Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Ala 115 120 125

Ile Ala Ala Tyr Leu Ala Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140

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Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

Arg Gly Ala Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

Ile Leu His Tyr Leu Lys Ala Ala Ala Tyr Ser His Cys Ala Trp Thr 130 135 140

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Thr Gly Tyr Leu Arg Asn 165

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Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

Arg Gly Ala Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ala Ala Cys Ala Trp Thr 130 135 140

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Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln
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Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln
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Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 . 80

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His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

Arg Gly Ala Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

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Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

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Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140

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Thr Gly Tyr Leu Arg Asn 165

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Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125 Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140

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Ala Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140

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Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140

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His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

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Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Ala 50 55 60

Asn Ile Ala Ser Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr
100 105 110

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140

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- Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45
- Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln
 50 55 60
- Asn Ile Phe Ala Ile Phe Ala Ala Ala Ser Ser Ser Thr Gly Trp Asn 65 70 75 80
- Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95
- His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110
- Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125
- Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140
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- Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln
 35 40 45
- Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln
 50 55 60
- Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80
- Ala Ser Ile Val Ala Ala Leu Leu Ser Asn Val Tyr His Gln Ile Asn 85 90 95
- His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr
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- Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125
- Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140

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Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

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Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

- Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60
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- His Leu Lys Thr Val Leu Ala Ala Lys Leu Ala Ala Ala Asp Phe Thr
- Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125
- Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140.
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- Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45
- Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60
- Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80
- Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95
- His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Ala Ala Thr 100 105 110
- Ala Gly Lys Ala Met Ser Ala Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125
- Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr 130 135 140
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Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

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Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

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Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60 Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120

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Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln 35 40 45

Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln 50 55 60

Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn 65 70 75 80

Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn 85 90 95

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr 100 105 110

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg 115 120 125

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ala Ala Cys Ala Trp Thr 130 135 140

Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Phe Ile Asn Arg Leu 145 150 155 160

Thr Gly Tyr Leu Arg Asn

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Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn

His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr

Arg Gly Lys Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg

Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr

Ile Val Arg Ala Glu Ile Leu Ala Asn Phe Ala Phe Ile Ala Arg Leu 155

Thr Gly Tyr Leu Arg Asn 165

Exhibit B

RAW SEQUENCE LISTING

The Biotechnology Systems Branch of the Scientific and Technical Information Center (STIC) no errors detected.

Application Serial Number:

Source:

Date Processed by STIC:

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IFW16

RAW SEQUENCE LISTING DATE: 08/25/2006 PATENT APPLICATION: US/10/802,540A TIME: 12:04:38

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Output Set: N:\CRF4\08252006\J802540A.raw

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             Whitty, Adrian
             Hochman, Paula
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      7 <130> FILE REFERENCE: BII-008.02
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70 <213> ORGANISM: Homo sapiens
71 <400> SEQUENCE: 3
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74 <210> SEQ ID NO: 4
75 <211> LENGTH: 39
76 <212> TYPE: DNA
77 <213> ORGANISM: Homo sapiens
78 <400> SEQUENCE: 4
79
         gccgctcgag ttatcagttt cggaggtaac ctgtaagtc
                                                                                  39
81 <210> SEQ ID NO: 5
82 <211> LENGTH: 35
83 <212> TYPE: DNA
84 <213> ORGANISM: Homo sapiens
85 <400> SEQUENCE: 5
        agetteeggg ggeeateate ateateatea taget
                                                                                  35
88 <210> SEQ ID NO: 6
89 <211> LENGTH: 35
90 <212> TYPE: DNA
91 <213> ORGANISM: Homo sapiens
92 <400> SEQUENCE: 6
93
        ccggagctat gatgatgatg atgatggccc ccgga
                                                                                 35
95 <210> SEQ ID NO: 7
96 <211> LENGTH: 87
97 <212> TYPE: DNA
98 <213> ORGANISM: Homo sapiens
99 <400> SEQUENCE: 7
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A

RAW SEQUENCE LISTING DATE: 08/25/2006
PATENT APPLICATION: US/10/802,540A TIME: 12:04:38

Input Set: N:\SSLM\J802540.raw
Output Set: N:\CRF4\08252006\J802540A.raw

100 101 103		ccggagacga tgatgacaag atggcttacg ccgctcttgg agccctacaa gcttctagca attttcagtg tcagaagctc ctgtggc SEO ID NO: 8	60 87
		LENGTH: 60	
		TYPE: DNA	
106	<213>	ORGANISM: Homo sapiens	
	<400>	SEQUENCE: 8	
108		gatctagcaa tgctgcctgt gctgccctcc tggctgcctt gaatgggagg cttgaatact	60
		SEG ID NO: A	
		LENGTH: 52 TYPE: DNA	
		ORGANISM: Homo sapiens	
		SEQUENCE: 9	
115		gcctcaagga caggatgaac tttgacatcc ctgaggagat taagcagctg ca	
117	<210>	SEQ ID NO: 10	52
		LENGTH: 76	
		TYPE: DNA	
120	<213>	ORGANISM: Homo sapiens	
121	<400>	SEQUENCE: 10	
123		aattgaatgg gagggctgca gcttgcgctg cagacaggat gaactttgac atccctgagg	60
	<210>	agattaagca gctgca SEQ ID NO: 11	76
		LENGTH: 76	
127	<212>	TYPE: DNA	
128	<213>	ORGANISM: Homo sapiens	
129	<400>	SEQUENCE: 11	
130		aattgaatgg gaggettgaa tactgeetea aggacaggge tgeatttget atceetgeag	60
131	-210-	agattaagca gctgca	76
		SEQ ID NO: 12 LENGTH: 51	
		TYPE: DNA	
		ORGANISM: Homo sapiens	
137	<400>	SEQUENCE: 12	
138		aattgaatgg gaggettgaa taetgeetca aggacaggat gaaetttgae a	51
	<210>	SEQ ID NO: 13	
		LENGTH: 43	
142	<212>	TYPE: DNA ORGANISM: Homo sapiens	
144	<400>	SEQUENCE: 13	
145		tccctgagga gattgctgca gctgcagctt tcgctgcagc tga	4.3
147	<210>	SEQ ID NO: 14	43
		LENGTH: 78	
149	<212>	TYPE: DNA	
150 •	<213>	ORGANISM: Homo sapiens	
151		SEQUENCE: 14	
152		egeogegettg accatetatg agatgetege taacateget agcattttca gacaagatte	60
	210>	atctagcact ggctggaa SEQ ID NO: 15	78
		LENGTH: 78	

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RAW SEQUENCE LISTING DATE: 08/25/2006 PATENT APPLICATION: US/10/802,540A TIME: 12:04:38

Input Set : N:\SSLM\J802540.raw

Output Set: N:\CRF4\08252006\J802540A.raw

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 158 <213> ORGANISM: Homo sapiens
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 160
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 161
            atctagcact ggctggaa
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 163 <210> SEQ ID NO: 16
 164 <211> LENGTH: 72
 165 <212> TYPE: DNA
 166 <213> ORGANISM: Homo sapiens
 167 <400> SEQUENCE: 16
 168
           ggaatgcttc aattgttgct gcactcctga gcaatgtcta tcatcagata aaccatctga
                                                                                    60
 169
           agacagttct ag
                                                                                    72
 171 <210> SEQ ID NO: 17
 172 <211> LENGTH: 72
 173 <212> TYPE: DNA
 174 <213> ORGANISM: Homo sapiens
 175 <400> SEQUENCE: 17
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 177
           ctgcagttct ag
                                                                                    72
 179 <210> SEQ ID NO: 18
 180 <211> LENGTH: 44
 181 <212> TYPE: DNA
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 182 <213> ORGANISM: Homo sapiens
 183 <400> SEQUENCE: 18
184
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186 <210> SEQ ID NO: 19
187 <211> LENGTH: 69
188 <212> TYPE: DNA
189 <213> ORGANISM: Homo sapiens
190 <400> SEQUENCE: 19
191
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                                                                                    60
192
          ctgaaaaga
                                                                                    69
194 <210> SEQ ID NO: 20
195 <211> LENGTH: 51
196 <212> TYPE: DNA
197 <213> ORGANISM: Homo sapiens
198 <400> SEQUENCE: 20
199
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                                                                                   51
201 <210> SEQ ID NO: 21
202 <211> LENGTH: 76
203 <212> TYPE: DNA
204 <213> ORGANISM: Homo sapiens
205 <400> SEQUENCE: 21
206
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                                                                                   60
207
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209 <210> SEQ ID NO: 22
210 <211> LENGTH: 87
211 <212> TYPE: DNA
212 <213> ORGANISM: Homo sapiens
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RAW SEQUENCE LISTING DATE: 08/25/2006
PATENT APPLICATION: US/10/802,540A TIME: 12:04:38

Input Set : N:\SSLM\J802540.raw

Output Set: N:\CRF4\08252006\J802540A.raw

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  214
  215
                                                                                     60
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                                                                                     87
  218 <211> LENGTH: 87
  219 <212> TYPE: DNA
  220 <213> ORGANISM: Homo sapiens
  221 <400> SEQUENCE: 23
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 222
 223
            ggagtacgct gcatgtgcct ggacgat
                                                                                     60
 225 <210> SEQ ID NO: 24
 226 <211> LENGTH: 50
 227 <212> TYPE: DNA
 228 <213> ORGANISM: Homo sapiens
 229 <400> SEQUENCE: 24
 230
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 232 <210> SEQ ID NO: 25
                                                                                    50
 233 <211> LENGTH: 166
 234 <212> TYPE: PRT
 235 <213> ORGANISM: Homo sapiens
 236 <400> SEQUENCE: 25
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 237
 238
                                                10
 239
           Cys Gln Lys Leu Leu Trp Gln Leu Asn Gly Arg Leu Glu Tyr Cys Leu
 240
                       20
           Lys Asp Arg Met Asn Phe Asp Ile Pro Glu Glu Ile Lys Gln Leu Gln
                                                                30
 241
 242
           Gln Phe Gln Lys Glu Asp Ala Ala Leu Thr Ile Tyr Glu Met Leu Gln
 243
 244
                                                        60
 245
           Asn Ile Phe Ala Ile Phe Arg Gln Asp Ser Ser Ser Thr Gly Trp Asn
246
                               70
247
           Glu Thr Ile Val Glu Asn Leu Leu Ala Asn Val Tyr His Gln Ile Asn
248
                           85
          His Leu Lys Thr Val Leu Glu Glu Lys Leu Glu Lys Glu Asp Phe Thr
249
250
                       100
                                           105
          Arg Gly Ala Leu Met Ser Ser Leu His Leu Lys Arg Tyr Tyr Gly Arg
251
252
253
          Ile Leu His Tyr Leu Lys Ala Lys Glu Tyr Ser His Cys Ala Trp Thr
                                                            125
254
                                   135
                                                       140
255
          Ile Val Arg Val Glu Ile Leu Arg Asn Phe Tyr Arg Ile Asn Arg Leu
256
                              150
                                                   155
257
          Thr Gly Tyr Leu Arg Asn
258
                          165
260 <210> SEQ ID NO: 26
261 <211> LENGTH: 166
262 <212> TYPE: PRT
263 <213 > ORGANISM: Homo sapiens
264 <400> SEQUENCE: 26
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VERIFICATION SUMMARY

PATENT APPLICATION: US/10/802,540A

DATE: 08/25/2006 TIME: 12:04:39

Input Set : N:\SSLM\J802540.raw
Output Set: N:\CRP4\08252006\J802540A.raw

L:9 M:271 C: Current Filing Date differs, Replaced Current Filing Date

STATISTICS SUMMARY

PATENT APPLICATION: US/10/802,540A

DATE: 08/25/2006 TIME: 12:04:39

Input Set : N:\SSLM\J802540.raw

Output Set: N:\CRF4\08252006\J802540A.raw

Application Serial Number: US/10/802,540A

Alpha or Numeric or Xml: Numeric

Application Class:

Application File Date: 03-26-2004

Art Unit: IFW16

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Software Application: PatentIN3.2

Total Number of Sequences: 56

Total Nucleotides: 1927

Total Amino Acids: 5495

Number of Brrors: 0

Number of Warnings: 0 Number of Corrections: 1

MESSAGE SUMMARY

271 C: 1 (Current Filing Date differs)